

CLAIMS

What is claimed is

1. A transgenic non-human mammalian animal having integrated within its genome a transgene encoding an exogenous wild-type α_{1A} , α_{1B} , or α_{1D} adrenergic receptor or a transgene encoding a constitutively-active mutant α_{1A} , α_{1B} , or α_{1D} adrenergic receptor, wherein the transgene is operably linked to a promoter that drives expression of the transgene in cells innervated by the sympathetic nervous system, and wherein the transgenic animal exhibits an abnormal phenotype.
2. The transgenic animal of claim 1 wherein the transgene encodes an exogenous wild-type α_{1B} adrenergic receptor or a constitutively active mutant α_{1B} adrenergic receptor.
3. The transgenic animal of claim 1 wherein the animal is a mouse and exhibits a neurodegenerative disorder-type phenotype.
4. The transgenic animal of claim 1 wherein the animal is a mouse and exhibits a phenotype resembling a cardiovascular disease.
5. The transgenic animal of claim 1 wherein the promoter is the promoter of the animal's endogenous α_{1B} adrenergic receptor.
6. The transgenic animals of claim 1 wherein the transgene encodes a constitutively active mutant hamster, rat, or human α_{1B} adrenergic receptor..
7. The transgenic animal of claim 1 wherein expression of the transgene results in the animal exhibiting Parkinson's disorder type symptoms..
8. The transgenic of animal of claim 1 wherein the transgene encoding a signal peptide.

9. A method of screening for a compound which modulates function of α_{1B} adrenergic receptor comprising:

administering the compound to the transgenic animal of claim 1; and

assaying for changes in the abnormal phenotype of said animal.

10. The method of claim 9 wherein the animal exhibits neurodegenerative symptoms and wherein the assay involves assaying for an improvement in or a delay in progression of the symptoms.

11. The method of claim 9 wherein the animal exhibits symptoms of a cardiovascular disorder and wherein the assay involves assaying for an improvement in or delay in progression of the symptoms.

12. The method of claim 9 wherein the assay involves evaluating the locomotor activity of the animal.

13. The method of claim 9 wherein the animal exhibits seizure type symptoms and wherein said assay involves evaluating the effect of the compound on the frequency, severity, or duration of said seizures.

14. A method of screening a drug for activity against a neurodegenerative disorder or a cardiovascular disorder, comprising

administering the drug to a transgenic mouse whose somatic cells comprise a transgene encoding an exogenous wild-type α_{1B} adrenergic receptor or a transgene encoding a constitutively-active mutant α_{1B} adrenergic receptor, wherein the transgene is operably linked to a promoter that drives expression of the transgene in cells innervated by the sympathetic nervous system, and wherein the transgenic animal exhibits symptoms characteristic of a disorder selected from the group consisting of a neurodegenerative disorder, a cardiovascular disorder, and a combination of a neurodegenerative and a cardiovascular disorder; and

monitoring the mouse for the effects of said drug on said symptoms.

15. The method of claim 14 wherein the transgenic mouse overexpresses an exogenous α_{1B} adrenergic receptor on the surface of cells in the brain of said animal.
- 5 16. The method of claim 14 wherein the transgenic mouse expresses a constitutively active mutant α_{1B} adrenergic receptor on the surface of cells in the brain of said animal.
17. A method for treating a subject with a neurodegenerative disorder, comprising:
administering to said subject a biologically effective amount of a compound
10 capable of blocking activation of α_1 adrenergic receptors
18. The method of claim 17 wherein said compound is an α_{1B} adrenergic receptor antagonist.
- 15 19. The method of claim 17 wherein said subject has exhibited symptoms characteristic of Parkinson's disease.
20. The method of claim 17 wherein the subject has exhibited seizures.
- 20 21. The method of claim 17 wherein the subject has exhibited locomotor impairment.